



United States
Environmental Protection Agency

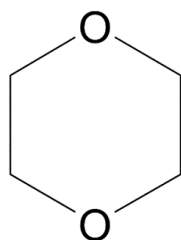
Office of Chemical Safety and
Pollution Prevention

Risk Evaluation for 1,4-Dioxane

Systematic Review Supplemental File:

Data Quality Evaluation of Environmental Hazard Studies

CASRN: 123-91-1



June 2019

Table of Contents

Table No.	Hero ID.	End-point(s)	Reference	Page Number
1	18670	Aquatic Vertebrates	Dawson, G. W., Jennings, A. L., Drozdowski, D., Rider, E. The acute toxicity of 47 industrial chemicals to fresh and saltwater fishes. <i>Journal of Hazardous Materials</i> . 1977.	1
2	3616460	Aquatic Invertebrates	G. Bringmann, R. Kuehn. 1982. Results of Toxic Action of Water Pollutants on <i>Daphnia magna</i> Straus Tested by an Improved Standardized Procedure. 15:1-6(GER) (ENG ABS) (OECDG Data File)	3
23	51735	Aquatic Plants	G. Bringmann, R. Kuhn. 1978. Grenzwerte der Schadwirkung wassergefährdender Stoffe gegen Blaualgen (<i>Microcystis aeruginosa</i>) und Grünalgen (<i>Scenedesmus quadricauda</i>) im Zellvermehrungshemmtest [Limiting values for the noxious effects of water pollutant material to blue algae (<i>Microcystis aeruginosa</i>) and green algae (<i>Scenedesmus quadricauda</i>) in cell propagation inhibition tests]. <i>Vom Wasser</i> 50:45-60	5
4	73652	Aquatic Invertebrates	G. Bringmann, R. Kuhn. 1977. The effects of water pollutants on <i>Daphnia magna</i> . <i>Wasser und Abwasser in Forschung und Praxis</i> 10:161-166	7
5	3616460	Aquatic Invertebrates	G. Bringmann, R. Kuehn. 1982. Results of Toxic Action of Water Pollutants on <i>Daphnia magna</i> Straus Tested by an Improved Standardized Procedure. 15:1-6(GER) (ENG ABS) (OECDG Data File)	9
6	3634436	Aquatic Invertebrates and Vertebrates	Brooke, L. 1987. Report of the Flow-Through and Static Acute Test Comparisons with Fathead Minnows and Acute Tests with Amphipod and a Cladoceran.	11
7	3660853	Aquatic Vertebrates	Geiger, D. L., Brooke, L. T., Call, D. J. 1990. Acute toxicities of organic chemicals to fathead minnows (<i>Pimephales promelas</i>): Volume V.	13
8	3661129	Aquatic Vertebrates	R. Johnson, J. Tietge, G. Stokes, D. Lothenbach. 1993. The Medaka Carcinogenesis Model.	15
9	4158026	Aquatic Vertebrates	Dow Chemical. 1989. 1,4-Dioxane: Embryo-larval toxicity test with the Fathead minnow, <i>Pimephales promelas</i> Rauschenque.	17
12	4438934	Aquatic Plants	G. Bringman, R. Kuhn. 1977. Limiting values of the harmful action of water endangering substances on bacteria (<i>Pseudomonas putida</i>) and green algae (<i>Scenedesmus quadricauda</i>) in the cell multiplication inhibition test. <i>Zeitschrift fuer Wasser- und Abwasser-Forschung</i> 10:87-98	19

Study Citation: Dawson, G. W., Jennings, A. L., Drozdowski, D., Rider, E.. 1977. The acute toxicity of 47 industrial chemicals to fresh and saltwater fishes. Journal of Hazardous Materials 1:303-318
 Data Type: Acute (0-96 hour); Aquatic; Fish
 Hero ID: 18670

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
	Metric 1: Test Substance Identity	High	× 2	2	
	Metric 2: Test Substance Source	Low	× 1	3	Test source was not reported.
	Metric 3: Test Substance Purity	Low	× 1	3	The test purity was not reported.
Domain 2: Test Design					
	Metric 4: Negative Controls	High	× 2	2	
	Metric 5: Negative Control Response	High	× 1	1	
	Metric 6: Randomized Allocation	Medium	× 1	2	Not specified, the information was implied for this metric.
Domain 3: Exposure Characterization					
	Metric 7: Experimental System/Test Media Preparation	Low	× 2	6	limited information provided
	Metric 8: Consistency of Exposure Administration	High	× 1	1	
	Metric 9: Measurement of Test Substance Concentration	High	× 1	1	
	Metric 10: Exposure Duration and Frequency	High	× 2	2	Information was provided for this metric.
	Metric 11: Number of Exposure Groups/Spacing of Exposure Levels	Low	× 1	3	No information was reported; however, an LC50 was derived.
	Metric 12: Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
	Metric 13: Test Organism Characteristics	High	× 2	2	
	Metric 14: Acclimitization and Pretreatment Conditions	High	× 1	1	
	Metric 15: Number of Organisms and Replicates per Group	Medium	× 1	2	Limited info provided
	Metric 16: Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
	Metric 17: Outcome Assessment Methodology	High	× 2	2	
	Metric 18: Consistency of Outcome Assessment	High	× 1	1	

Continued on next page ...

... continued from previous page

Study Citation: Dawson, G. W., Jennings, A. L., Drozdowski, D., Rider, E.. 1977. The acute toxicity of 47 industrial chemicals to fresh and saltwater fishes. Journal of Hazardous Materials 1:303-318
 Data Type: Acute (0-96 hour); Aquatic; Fish
 Hero ID: 18670

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 6: Confounding / Variable Control					
Metric 19:	Confounding Variables in Test Design and Procedures	High	× 2	2	Information was reported for this metric.
Metric 20:	Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
Metric 21:	Statistical Methods	High	× 1	1	
Metric 22:	Reporting of Data	High	× 2	2	
Metric 23:	Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.4	
Extracted		No			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.

Study Citation:	G. Bringmann, R. Kuehn. 1982. Ergebnisse der Schadwirkung wassergefährdender Stoffe gegen Daphnia magna in einem weiterentwickelten standardisierten Testverfahren [Results of toxic action of water pollutants on Daphnia magna Straus tested by an improved standardized procedure]. Wasser und Abwasser in Forschung und Praxis 15				
Data Type:	Acute (0-96 hour); Aquatic; Invertebrates				
Hero ID:	18804				

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	Low	× 2	6	Only the chemical name was provided.
Metric 2:	Test Substance Source	Low	× 1	3	The source of the chemical was not reported
Metric 3:	Test Substance Purity	Low	× 1	3	The information for this metric was provided in the report.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	The information for this metric was provided in the report.
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	

Continued on next page ...

... continued from previous page

Study Citation:	G. Bringmann, R. Kuehn. 1982. Ergebnisse der Schadwirkung wassergefährdender Stoffe gegen Daphnia magna in einem weiterentwickelten standardisierten Testverfahren [Results of toxic action of water pollutants on Daphnia magna Straus tested by an improved standardized procedure]. Wasser und Abwasser in Forschung und Praxis 15				
Data Type:	Acute (0-96 hour); Aquatic; Invertebrates				
Hero ID:	18804				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
	Metric 18: Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	N/A		N/A	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	Medium	× 1	2	Limited information was provided for this metric.
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.3	
Extracted		Yes			

* MWF = Metric Weighting Factor

† High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

‡ The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left[\frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right]_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

†† This metric met the criteria for high confidence as expected for this type of study.

Study Citation: G. Bringmann, R. Kuhn. 1978. Grenzwerte der Schadwirkung wassergefährdender Stoffe gegen Blaualgen (*Microcystis aeruginosa*) und Grünalgen (*Scenedesmus quadricauda*) im Zellvermehrungshemmtest [Limiting values for the noxious effects of water pollutant material to blue algae (*Microcystis aeruginosa*) and green algae (*Scenedesmus quadricauda*) in cell propagation inhibition tests].. Vom Wasser 50:45-60

Data Type: Other; Aquatic; Plants

Hero ID: 51735

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	Low	× 2	6	Only the chemical name was provided.
Metric 2:	Test Substance Source	Low	× 1	3	The source of the chemical was not reported.
Metric 3:	Test Substance Purity	Low	× 1	3	Information about the test purity was not reported.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	N/A		N/A	The solubility for 1,4-dioxane was not a factor.
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimatization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	N/A		N/A	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	

Continued on next page . . .

... continued from previous page

Study Citation: G. Bringmann, R. Kuhn. 1978. Grenzwerte der Schadwirkung wassergefährdender Stoffe gegen Blaualgen (*Microcystis aeruginosa*) und Grünalgen (*Scenedesmus quadricauda*) im Zellvermehrungshemmtest [Limiting values for the noxious effects of water pollutant material to blue algae (*Microcystis aeruginosa*) and green algae (*Scenedesmus quadricauda*) in cell propagation inhibition tests].. Vom Wasser 50:45-60
 Data Type: Other; Aquatic; Plants
 Hero ID: 51735

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
	Metric 18: Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	Medium	× 1	2	Limited information was provided for this metric.
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.3	
Extracted		Yes			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.

Study Citation: G. Bringmann, R. Kuhn. 1977. The effects of water pollutants on *Daphnia magna*. Wasser und Abwasser in Forschung und Praxis 10:161-166
 Data Type: Acute (0-96 hour); Aquatic; Invertebrates
 Hero ID: 73652

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	Low	× 2	6	No chemical identity information was provided for any of the chemicals including 1,4-Dioxane in this report. Only the chemical names were provided.
Metric 2:	Test Substance Source	Low	× 1	3	The report source did not provide any information about the manufacturer of the chemicals tested.
Metric 3:	Test Substance Purity	Low	× 1	3	Information about the purity was not provided.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	Low	× 1	3	No information was provided for this metric.
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					

Continued on next page ...

... continued from previous page

Study Citation:	G. Bringmann, R. Kuhn. 1977. The effects of water pollutants on Daphnia magna. Wasser und Abwasser in Forschung und Praxis 10:161-166				
Data Type:	Acute (0-96 hour); Aquatic; Invertebrates				
Hero ID:	73652				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
	Metric 17: Outcome Assessment Methodology	High	× 2	2	This metric was completely characterized.
	Metric 18: Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	Low	× 1	3	No statistical methods were provided.
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.4	
Extracted		Yes			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.

Study Citation: G. Bringmann, R. Kuehn. 1982. Results of Toxic Action of Water Pollutants on Daphnia magna Straus Tested by an Improved Standardized Procedure. 15:1-6(GER) (ENG ABS) (OECDG Data File)
 Data Type: Acute (0-96 hour); Aquatic; Invertebrates
 Hero ID: 3616460

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	Low	× 2	6	The study only listed the test substance name. However, this reporting source was for multiple chemicals thus detail information could have been omitted because of publication requirements. Also, this study was published in 1983 from Germany.
Metric 2:	Test Substance Source	Low	× 1	3	The study only listed the test substance name. However, this reporting source was for multiple chemicals thus detail information could have been omitted because of publication requirements. Also, this study was published in 1983 from Germany.
Metric 3:	Test Substance Purity	Low	× 1	3	The study only listed the test substance name. However, this reporting source was for multiple chemicals thus detail information could have been omitted because of publication requirements. Also, this study was published in 1983 from Germany.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	N/A		N/A	This information was reported for this metric.
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	N/A		N/A	

Continued on next page ...

... continued from previous page

Study Citation:	G. Bringmann, R. Kuehn. 1982. Results of Toxic Action of Water Pollutants on Daphnia magna Straus Tested by an Improved Standardized Procedure. 15:1-6(GER) (ENG ABS) (OECDG Data File)					
Data Type:	Acute (0-96 hour); Aquatic; Invertebrates					
Hero ID:	3616460					
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}	
	Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
	Metric 15:	Number of Organisms and Replicates per Group	N/A		N/A	
	Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment						
	Metric 17:	Outcome Assessment Methodology	High	× 2	2	
	Metric 18:	Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control						
	Metric 19:	Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20:	Outcomes Unrelated to Exposure	High	× 1	1	This information was reported for this metric.
Domain 7: Data Presentation and Analysis						
	Metric 21:	Statistical Methods	Low	× 1	3	Limited information was provided for this metric. Only effects results were reported.
	Metric 22:	Reporting of Data	High	× 2	2	
	Metric 23:	Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]			High		1.4	
Extracted			Yes			

* MWF = Metric Weighting Factor

† High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

‡ The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

†† This metric met the criteria for high confidence as expected for this type of study.

Study Citation: Brooke, L.. 1987. Report of the Flow-Through and Static Acute Test Comparisons with Fathead Minnows and Acute Tests with an Amphipod and a Cladoceran.
 Data Type: Acute (0-96 hour); Aquatic; other Fish and Daphnia
 Hero ID: 3634436

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	High	× 2	2	
Metric 2:	Test Substance Source	High	× 1	1	
Metric 3:	Test Substance Purity	High	× 1	1	
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	
Metric 18:	Consistency of Outcome Assessment	High	× 1	1	

Continued on next page ...

... continued from previous page

Study Citation:	Brooke, L.. 1987. Report of the Flow-Through and Static Acute Test Comparisons with Fathead Minnows and Acute Tests with an Amphipod and a Cladoceran.				
Data Type:	Acute (0-96 hour); Aquatic; other Fish and Daphnia				
Hero ID:	3634436				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	High	× 1	1	
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.0	
Extracted		Yes			

* MWF = Metric Weighting Factor

† High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

‡ The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

†† This metric met the criteria for high confidence as expected for this type of study.

Study Citation: Geiger, D. L., Brooke, L. T., Call, D. J.. 1990. Acute toxicities of organic chemicals to fathead minnows (*Pimephales promelas*): Volume V.
 Data Type: Acute (0-96 hour); Aquatic; Fish
 Hero ID: 3660853

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	High	× 2	2	
Metric 2:	Test Substance Source	High	× 1	1	
Metric 3:	Test Substance Purity	High	× 1	1	
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimatization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	
Metric 18:	Consistency of Outcome Assessment	High	× 1	1	

Continued on next page ...

... continued from previous page

Study Citation:	Geiger, D. L., Brooke, L. T., Call, D. J.. 1990. Acute toxicities of organic chemicals to fathead minnows (<i>Pimephales promelas</i>): Volume V.				
Data Type:	Acute (0-96 hour); Aquatic; Fish				
Hero ID:	3660853				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	High	× 1	1	
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.0	
Extracted		Yes			

* MWF = Metric Weighting Factor

† High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

‡ The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lfloor \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rfloor_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases},$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

†† This metric met the criteria for high confidence as expected for this type of study.

Study Citation: R. Johnson, J. Tietge, G. Stokes, D. Lothenbach. 1993. The Medaka Carcinogenesis Model.
 Data Type: Chronic (>21 days); Aquatic; Fish
 Hero ID: 3661129

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	High	× 2	2	The chemical identity was provided in this study.
Metric 2:	Test Substance Source	Low	× 1	3	The manufacturer was not provided.
Metric 3:	Test Substance Purity	Low	× 1	3	The purity was not provided for this study.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	Low	× 1	3	Other chemicals were tested. This could have affected the choice for not providing this information.
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	Medium	× 1	2	Limited information was provided.
Metric 12:	Testing at or Below Solubility Limit	N/A		N/A	This test substance is very soluble.
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	
Metric 18:	Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control					

Continued on next page ...

... continued from previous page

Study Citation:	R. Johnson, J. Tietge, G. Stokes, D. Lothenbach. 1993. The Medaka Carcinogenesis Model.				
Data Type:	Chronic (>21 days); Aquatic; Fish				
Hero ID:	3661129				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	This endpoint is well characterized.
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	High	× 1	1	
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.2	
Extracted		Yes			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left[\sum_i (\text{Metric Score}_i \times \text{MWF}_i) / \sum_j \text{MWF}_j \right]_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.

Study Citation: Dow Chemical. 1989. 1,4-Dioxane: Embryo-larval toxicity test with the Fathead minnow, *Pimephales promelas* Rafinesque.
 Data Type: Acute (0-96 hour); Aquatic; Fish
 Hero ID: 4158026

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	High	× 2	2	
Metric 2:	Test Substance Source	High	× 1	1	
Metric 3:	Test Substance Purity	High	× 1	1	
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	The information was provided for this metric.
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimitization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	The information was provided for this metric.
Metric 18:	Consistency of Outcome Assessment	High	× 1	1	
Domain 6: Confounding / Variable Control					
Continued on next page ...					

... continued from previous page

Study Citation:	Dow Chemical. 1989. 1,4-Dioxane: Embryo-larval toxicity test with the Fathead minnow, Pimephales promelas Rafinesque.				
Data Type:	Acute (0-96 hour); Aquatic; Fish				
Hero ID:	4158026				
Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	High	× 1	1	
	Metric 22: Reporting of Data	High	× 2	2	
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.0	
Extracted		Yes			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left[\sum_i (\text{Metric Score}_i \times \text{MWF}_i) / \sum_j \text{MWF}_j \right]_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.

Study Citation: G. Bringman, R. Kuhn. 1977. Limiting values of the harmful action of water endangering substances on bacteria (*Pseudomonas putida*) and green algae (*Scenedesmus quadricauda*) in the cell multiplication inhibition test. *Zeitschrift fuer Wasser- und Abwasser-Forschung* 10:87-98

Data Type: Other; Aquatic; Plants

Hero ID: 4438934

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 1: Test Substance					
Metric 1:	Test Substance Identity	Low	× 2	6	Only the chemical name was provided for this study.
Metric 2:	Test Substance Source	Low	× 1	3	The source was not reported.
Metric 3:	Test Substance Purity	Low	× 1	3	The test purity was not reported.
Domain 2: Test Design					
Metric 4:	Negative Controls	High	× 2	2	
Metric 5:	Negative Control Response	High	× 1	1	
Metric 6:	Randomized Allocation	High	× 1	1	
Domain 3: Exposure Characterization					
Metric 7:	Experimental System/Test Media Preparation	High	× 2	2	
Metric 8:	Consistency of Exposure Administration	High	× 1	1	
Metric 9:	Measurement of Test Substance Concentration	High	× 1	1	
Metric 10:	Exposure Duration and Frequency	High	× 2	2	
Metric 11:	Number of Exposure Groups/Spacing of Exposure Levels	High	× 1	1	
Metric 12:	Testing at or Below Solubility Limit	High	× 1	1	
Domain 4: Test Organism					
Metric 13:	Test Organism Characteristics	High	× 2	2	
Metric 14:	Acclimatization and Pretreatment Conditions	High	× 1	1	
Metric 15:	Number of Organisms and Replicates per Group	High	× 1	1	
Metric 16:	Adequacy of Test Conditions	High	× 1	1	
Domain 5: Outcome Assessment					
Metric 17:	Outcome Assessment Methodology	High	× 2	2	
Metric 18:	Consistency of Outcome Assessment	High	× 1	1	

Continued on next page . . .

... continued from previous page

Study Citation: G. Bringman, R. Kuhn. 1977. Limiting values of the harmful action of water endangering substances on bacteria (*Pseudomonas putida*) and green algae (*Scenedesmus quadricauda*) in the cell multiplication inhibition test. *Zeitschrift fuer Wasser- und Abwasser-Forschung* 10:87-98

Data Type: Other; Aquatic; Plants

Hero ID: 4438934

Domain	Metric	Rating [†]	MWF*	Score	Comments ^{††}
Domain 6: Confounding / Variable Control					
	Metric 19: Confounding Variables in Test Design and Procedures	High	× 2	2	
	Metric 20: Outcomes Unrelated to Exposure	High	× 1	1	
Domain 7: Data Presentation and Analysis					
	Metric 21: Statistical Methods	High	× 1	1	
	Metric 22: Reporting of Data	Medium	× 2	4	Limited information was provided for this metric.
	Metric 23: Explanation of Unexpected Outcomes	N/A		N/A	
Overall Quality Determination [‡]		High		1.3	
Extracted		Yes			

* MWF = Metric Weighting Factor

[†] High = 1; Medium = 2; Low = 3; Unacceptable = 4; N/A has no value.

[‡] The overall rating is calculated as necessary. EPA may not always provide a comment for a metric that has been categorized as High.

$$\text{Overall rating} = \begin{cases} 4 & \text{if any metric is Unacceptable} \\ \left\lceil \frac{\sum_i (\text{Metric Score}_i \times \text{MWF}_i)}{\sum_j \text{MWF}_j} \right\rceil_{0.1} & \text{(round to the nearest tenth) otherwise} \end{cases}$$

where High = ≥ 1 to < 1.7 ; Medium = ≥ 1.7 to < 2.3 ; Low = ≥ 2.3 to ≤ 3 . If the reviewer determines that the overall rating needs adjustment, the original rating is crossed out and an arrow points to the new rating.

^{††} This metric met the criteria for high confidence as expected for this type of study.